EXHIBIT I

Wendy Thai

Subject: FW: [help #21742] RE: GROUP:Q9H5V8

Attachments: Wendy Thai.vcf

-----Original Message-----
From: Elisabeth Gasteiger via RT [mailto:help@uniprot.org]
Sent: Wednesday, June 20, 2007 4:15 AM
To: Wendy Thai
Subject: [help #21742] RE: GROUP:Q9H5V8

> [WThai@slwk.com - Wed Jun 20 00:25:32 2007]:
>
> Hi Elisabeth:
>
> Thank you for your help with this.
>
> I was told by an examiner at the US Patent and Trademark Office that the
> Scherl-Mostageer sequence had been corrected.
>
> Also, my sequence analysis data (see attached WORD document) indicates
> that the Scherl-Mostageer sequence has been changed in the database.
>
> For example, when I do an alignment of the original Scherl-Mostageer
> sequence with my sequence (SIMA135), I see two amino acid mismatches -
> at positions 525 and 827. See page 16 of the attached WORD document.
>
> However, when I do a blastp analysis of my sequence (SIMA135) against
> the database, I get 100 % identity with Q9H5V8. See page 1 of the
> attached WORD document. Q9H5V8 references the Scherl-Mostageer sequence
> (reference 1) - SEE pages 7-8 of attached WORD document.
>
> Thus, it appears as if the Scherl-Mostageer sequence had been changed
> and is now identical to SIMA135. Please advise. Am I misreading the
> sequence analysis results? If you think a telephone call would be
> helpful, let me know and I can call you.
>
> Thanks so much for your help.
>
> Wendy Thai
> 612-373-6913

Dear Wendy,
Thank you for these precisions.
The 2 mismatches are indeed reported in the UniProtKB/Swiss-Prot record.
The first one is a known variant:
FT VARIANT 525 525 R -> Q (in dbSNP:rs3749191).
FT /FTId=VAR_025498.

And the second is annotated as a sequencing conflict (ref.1 is the
Scherl-Mostageer paper in the entry):
FT CONFLICT 827 827 S -> N (in Ref. 1, 3 and 5; BAB14695).

Your confusion is indeed caused by the fact that **Swiss-Prot is a non-redundant database that strives to have 1 protein entry for each gene product. When the sequence was manually annotated, the scientist who did the annotation saw that Q9H5V8 and Q96QU7 (the TrEMBL entry translated from the Scherl-Mostageer sequence AY026461) described the same protein although there sequences differed slightly (in 2 residues).**

And here is where the answer I gave you yesterday was not quite correct: I said the sequences were 100% identical but indeed they are not. Here is the entry history
http://www.ebi.ac.uk/uniprot/unisave/?query=Q9H5V8&search=Go

and here is the portion of our user manual which describes the concept of minimal redundancy:
http://www.expasy.org/sprot/userman.html#what_is_sprot

In summary, the Scherl-Mostageer sequence has not changed since its submission. It is just in the manually annotated Swiss-Prot database that it has been considered to describe the same protein as the TrEMBL sequence with which it was merged (note that this is an old version of the Q9H5V8 entry from before the merge - it was in TrEMBL then and moved to Swiss-Prot upon manual annotation):

```
ID   Q9H5V8_HUMAN   PRELIMINARY;   PRT;   836 AA.
AC   Q9H5V8;
DT   01-MAR-2001, integrated into UniProtKB/TrEMBL.
DT   01-MAR-2001, sequence version 1.
DT   07-FEB-2006, entry version 13.
DE   Hypothetical protein FLJ22969 (NCSG135).
OS   Homo sapiens (Human).
OC   Eukaryota; Metazoa; Chordata; Craniata; Vertebrata; Euteleostomi;
OC   Mammalia; Eutheria; Euarchontoglires; Primates; Catarrhini; Hominidae;
OC   Homo.
OX   NCBI_TaxID=9606;
RN   [1]
RP   NUCLEOTIDE SEQUENCE.
RA   Watanabe K., Kumagai A., Itakura S., Yamazaki M., Tashiro H., Ota T.,
RA   Suzuki Y., Obayashi M., Nishi T., Shibahara T., Tanaka T.,
RA   Nakamura Y., Isogai T., Sugano S.;
RL   Submitted (AUG-2000) to the EMBL/GenBank/DDBJ databases.
RN   [2]
RP   NUCLEOTIDE SEQUENCE.
RX   MEDLINE=22547370; PubMed=12660814; DOI=10.1038/sj.onc.1206220;
RA   Hooper J.D., Zijlstra A., Aimes R.T., Liang H., Claassen G.F.,
RA   Tarin D., Testa J.E., Quigley J.P.;
RT   "Subtractive immunization using highly metastatic human tumor cells
RT   identifies SIMA135/CDCP1, a 135 kDa cell surface phosphorylated
RT   glycoprotein antigen.";
RL   Oncogene 22:1783-1794(2003).
CC   ---------------------------------------------------------------------
CC   Copyrighted by the UniProt Consortium, see http://www.uniprot.org/terms
CC   Distributed under the Creative Commons Attribution-NoDerivs License
CC   ---------------------------------------------------------------------
DR   EMBL; AK026622; BAB15511.1; -; mRNA.
DR   EMBL; AF468010; AAO33397.1; -; mRNA.
DR   Ensembl; ENSG00000163814; Homo sapiens.
SQ   SEQUENCE   836 AA;  92875 MW;  9B980475C3E5C4C8 CRC64;
     MAGLNCGVSI ALLGVLLLGA ARLPRGAEAF EIALPRESNI TVLIKLGTPT LLAKPCYIVI
     SKRHITMLSI KSGERIVFTF SCQSPENHFV IEIQKNIDCM SGPCPFGEVQ LQPSTSLLPT
     LNRTFIWDVK AHKSIGLELQ FSIPRLRQIG PGESCPDGVT HSISGRIDAT VVRIGTFCSN
     GTVSRIKMQE GVKMALHLPW FHPRNVSGFS IANRSSIKRL CIIESVFEGE GSATLMSANY
     PEGFPEDELM TWQFVVPAHL RASVSFLNFN LSNCERKEER VEYYIPGSTT NPEVFKLEDK
     QPGNMAGNFN LSLQGCDQDA QSPGILRLQF QVLVQHPQNE SNKIYVVDLS NERAMSLTIE
     PRPVKQSRKF VPGCFVCLES RTCSSNLTLT SGSKHKISFL CDDLTRLWMN VEKTISCTDH
     RYCQRKSYSL QVPSDILHLP VELHDFSWKL LVPKDRLSLV LVPAQKLQQH THEKPCNTSF
     SYLVASAIPS QDLYFGSFCP GGSIKQIQVK QNISVTLRTF APSFRQEASR QGLTVSFIPY
     FKEEGVFTVT PDTKSKVYLR TPNWDRGLPS LTSVSWNISV PRDQVACLTF FKERSGVVCQ
     TGRAFMIIQE QRTRAEEIFS LDEDVLPKPS FHHHSFWVNI SNCSPTSGKQ LDLLFSVTLT
     PRTVDLTVIL IAAVGGGVLL LSALGLIICC VKKKKKKTNK GPAVGIYNGN INTEMPRQPK
     KFQKGRKDND SHVYAVIEDT MVYGHLLQDS SGSFLQPEVD TYRPFQGTMG VCPPSPPTIC
     SRAPTAKLAT EEPPPRSPPE SESEPYTFSH PNNGDVSSKD TDIPLLSTQE PMEPAE
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I hope this helps.

Best regards
Elisabeth Gasteiger

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Wendy Thai.vcf
(182 B)
